



DECLARATION AND POWER OF ATTORNEY

| (if only one name is listed below) o is claimed and for which a patent HUMAN PDGF BETA RECEPT Application Serial No. 08/258,283 | nd citizenship are as stated be r an original, first and joint in is sought on the invention en OR the specification of which and was amended on | ventor (if plural inventors are relatitled: INHIBITORY IMMU is attached hereto or (if applicable). | |
|---|--|---|--|
| referred to above. I acknowledge t with Title 37, Code of Federal Reg | he duty to disclose informational rulations, Section 1.56. I clai for patent or inventor's certific | n which is material to the exan m foreign priority benefits und cate listed below and have also | e claims, as amended by any amendment nination of this application in accordance ler Title 35, United States Code, Section identified below any foreign application priority is claimed. |
| Prior Foreign Application(s) | ά. | | |
| Country | Application No. | Date of Filing | Priority Claimed Under 35 USC 119 |
| | | · | Yes No |
| | | - | Yes No |

I claim the benefit under Title 35, United States Code, Section 120 of any United States application(s) listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States application in the manner provided by the first paragraph of Title 35, United States Code, Section 112, I acknowledge the duty to disclose material information as defined in Title 37, Code of Federal Regulations, Section 1.56 which occurred between the filing date of the prior application and the national or PCT international filing date of this application:

| Application Serial No. | Date of Filing | Status | |
|------------------------|------------------|------------------------------|--|
| Presently Unassigned | June 7, 1994 | PatentedX Pending Abandoned | |
| 07/801,795 | December 2, 1991 | Patented X Pending Abandoned | |

POWER OF ATTORNEY: As a named inventor, I hereby appoint the following attorney(s) and/or agent(s) to prosecute this application and transact all business in the Patent and Trademark Office connected therewith.

William B. Smith, Reg. No. 30,223 Karen B. Dow, Reg. No. 29,684 Randolph T. Apple, Reg. No. 36,429

| Send Correspondence to: Karen B. Dow TOWNSEND and TOWNSEND KHOURIE and CREW | Direct Telephone Calls to: (Name, Reg. No., telephone no.) |
|---|--|
| Steuart Street Tower | Karen B. Dow |
| One Market Plaza, 20th Floor | Reg. No. 29,684 |
| San Francisco, CA 94105 | (415) 326-2400, Ext. 217 |





| Full Name | Last Name | First Name | Middle Name or In | itial |
|----------------------------|---|----------------------------------|--|-------------------|
| of Inventor 1 | Ramakrishnan | Vanitha | | · |
| Residence & Citizenship | City Belmont | State/Foreign Country California | Country of Citizens India | ship |
| Post Office Address | Post Office Address 2019 Arbor Avenue | City Belmont | State/Country California | Zip Code 94002 |
| Full Name of Inventor 2 | Last Name Escobedo | First Name Maria | Middle Name or Initial Amelia | |
| Residence & Citizenship | City San Francisco | State/Foreign Country California | Country of Citizens Chile | hip |
| Post Office Address | Post Office Address 455 Upper Terrace #3 | City San Francisco | State/Country California | Zip Code 94117 |
| Full Name of Inventor 3 | Last Name Fretto | First Name Larry | Middle Name or Initial J. | |
| Residence & Citizenship | City Belmont | State/Foreign Country California | Country of Citizenship U.S.A. | |
| Post Office Address | Post Office Address 1553 Escondido Way | City Belmont | State/Country California | Zip Code 94002 |
| Full Name of Inventor 4 | Last Name Lokker | First Name Nathalie | Middle Name or Initial | |
| Residence & Citizenship | City San Francisco | State/Foreign Country California | Country of Citizenship The Netherlands | |
| Post Office Address | Post Office Address 741 40th Avenue | City San Francisco | State/Country California | Zip Code 94121 |
| Full Name of Inventor 5 | Last Name | First Name | Middle Name or Initial | |
| Residence & Citizenship | City | State/Foreign Country | Country of Citizenship | |
| Post Office Address | Post Office Address | City | State/Country | Zip Code |
| Full Name of Inventor 6 | Last Name | First Name | Middle Name or Initial | |
| Residence & Citizenship | City | State/Foreign Country | Country of Citizenship | |
| Post Office Address | Post Office Address | City | State/Country | Zip Code |

I further declare that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code, and that such willful false statements may jeopardize the validity of the application or any patent issuing thereon.

| Signature of Inventor 1 [autha/amaknshaan] | Signature of Inventor 2 | Signature of Inventor 3 |
|---|-------------------------|-------------------------|
| Vanitha Ramakrishnan | Maria Amelia Escobedo | Larry J. Fretto |
| Date 8 2 94 | Date \$2/94 | Date 8-2-94 |
| Signature of Inventor 4 Nathalie Lokke | Signature of Inventor 5 | Signature of Inventor 6 |
| Date 17 8/2/94 | Date | Date |

K:\12418\18-2DEC.POA

(Page 3 of 3)

VER DESTATEMENT (DECLARATION) CLAIMING SWALL ENTITY STATUS (37 CFR 1.9(f) & 1.27(c)) - SMALL BUSINESS CONCERN

| -,- | | , Maria Amelia Escobedo, Larry J. Fretto and | Nathalie Lokker |
|--|---|---|--|
| Serial or Patent I Filed or Issued: | | · | |
| | | DLYPEPTIDES TO HUMAN PDGF BETA RI | ECEPTOR |
| I hereby declare | | | |
| | • • | e small business concern identified below: e small business concern empowered to act on | behalf of the concern identified below: |
| NAME OF SM | ALL BUSINESS CONCERN | .COR THERAPEUTICS, INC. | |
| | SMALL BUSINESS CONCER | | CA 94080 |
| in 37 CFR 1.9(d) concern, includin concern is the ave of the pay period |), for purposes of paying reduction the control of | ced fees to the United States Patent and Trade tot exceed 500 persons. For purposes of this stay year of the concern of the persons employed on | oncern as defined in 13 CFR 121.12, and reproduced mark Office, in that the number of employees of the atement, (1) the number of employees of the business a full-time, part-time or temporary basis during each directly or indirectly, one concern controls or has the oth. |
| to the invention, | | UNOGLOBULIN POLYPEPTIDES TO HUMA | small business concern identified above with regard N PDGF BETA RECEPTOR by inventor(s) Vanitha |
| [] | the specification filed here | with | |
| [X] | application Serial No. | 08/258,283 , filed June | 10, 1994 |
| .[] | Patent No. | , issued | |
| *NOTE: Separat | e verified statements are requinities. (37 CFR 1.27) | | zation having rights to the invention averring to their |
| NAME | [] INDIVIDUAL | [] SMALL BUSINESS CONCERN | [] NONPROFIT ORGANIZATION |
| ADDRESS | [] INDIVIDUAL | [] SMALL BUSINESS CONCERN | [] NONPROFIT ORGANIZATION |
| prior to paying, o | | | resulting in loss of entitlement to small entity status fue after the date on which status as a small entity is |
| to be true; and fur or imprisonment, | rther that these statements wer or both, under section 1001 of | e made with the knowledge that willful false st | tements made on information and belief are believed attements and the like so made are punishable by fine th willful false statements may jeopardize the validity is directed. |
| NAME OF PER | SON SIGNING | P-200- 1 Silit | · |
| | SON IF OTHER THAN OWN | NER ROBER L SWIT | 1.61-1.59708017 |
| • / | PERSON/SIGNING | 256 E. Grand Avenue, South San Fra | ncisco, CA 94080 |
| SIGNATURE _ | (ll) | DATE | 8/2/14 |
| K:\12416\18-2.SES | | /) ` | |



| As a below named inventor, I declare that: |
|---|
| My residence, post office address and citizenship are as stated below next to my name; I believe I am the original, first and sole inventor |
| (if only one name is listed below) or an original, first and joint inventor (if plural inventors are named below) of the subject matter which |
| is claimed and for which a patent is sought on the invention entitled: INHIBITORY IMMUNOGLOBULIN POLYPEPTIDES TO |
| HUMAN PDGF HETA RECEPTOR the specification of which is attached hereto or X was filed on June 10, 1994 at |
| Application Serial No. 08/258,283 and was amended on (if applicable). |

I have reviewed and understand the contents of the above identified specification, including the claims, as amended by any amendment referred to above. I acknowledge the duty to disclose information which is material to the examination of this application in accordance with Title 37, Code of Federal Regulations, Section 1.56. I claim foreign priority benefits under Title 35, United States Code, Section 119 of any foreign applications(s) for patent or inventor's certificate listed below and have also identified below any foreign application for patent or inventor's certificate having a filing date before that of the application on which priority is claimed.

Prior Foreign Application(s)

| Country | Application No. | Date of Filing | Priority Claimed Under 35 USC 119 | |
|---------|-----------------|----------------|--------------------------------------|--|
| | | | Yes No | |
| | | | Yes No | |

I claim the benefit under Title 35. United States Code, Section 120 of any United States application(s) listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States application in the manner provided by the first paragraph of Title 35, United States Code, Section 112, I acknowledge the duty to disclose material information as defined in Title 37, Code of Federal Regulations, Section 1.56 which occurred between the filing date of the prior application and the national or PCT international filing date of this application:

| Application Serial No. | Date of Filing | Status | |
|------------------------|------------------|-------------------------------|--|
| 08/253,440 | June 7, 1994 | Patented X Pending Abandoned | |
| 07/801,795 | December 2, 1991 | Patented _X Pending Abandoned | |

POWER OF ATTORNEY: As a named inventor, I hereby appoint the following attorney(s) and/or agent(s) to prosecute this application and transact all business in the Patent and Trademark Office connected therewith.

Shelley Precivale Eberle, Rag. No. 31,411 Karen B. Dow, Reg. No. 29,684 Randolph T. Apple, Reg. No. 36,429

| Send Correspondence to: Karen B. Dow TOWNSEND and TOWNSEND KHOURIE and CREW | Direct Telephone Calls to: (Name, Reg. No., telephone 20.) |
|---|---|
| Steuart Street Tower | Karen B. Dow |
| One Market Plaza, 20th Floor San Francisco, CA 94105 | Reg. No. 29,684 (415) 326-2400, Ext. 217 |

| Full Name of Inventor 1 | Last Name Ramakrishnan | First Name Vaniths | Middle Name or Initial | |
|----------------------------|--|-------------------------------------|-------------------------------|----------|
| Residence & Citizenship | City Relmont | State/Foreign Country California | Country of Citizenship India | |
| Post Office | Post Office Address | Ciry | State/Country Zip Code | |
| Address | 2019 Arbor Avenue | Belmont | California 94002 | |
| Full Name | Last Name | First Name | Middle Name or Initial Amelia | |
| of Inventor 2 | Escobedo | Maria | | |
| Residence & | City | State/Foreign Country | Country of Citizenship Chile | |
| Citizenship | San Francisco | California | | |
| Post Office | Post Office Address 455 Upper Terrace #3 | City | State/Country | Zip Code |
| Address | | San Francisco | California | 94117 |
| Full Name of Inventor 3 | Last Name Fretto | First Name Larry | Middle Name or Initial J. | |
| Residence & Citizenship | City Belmont | State/Foreign Country California | Country of Citizenship U.S.A. | |
| Post Office | Post Office Address | City | State/Country | Zip Code |
| Address | 1553 Escondido Way | Belmont | California | 94002 |

I further declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code, and that such willful false statements may jeopardize the validity of the application or any patent issuing thereon.

| Signature of Inventor 1 | Signature of Inventor 2 Signature of Inventor 3 Cleaner Q. Esco Stelly | |
|-------------------------|--|-----------------|
| Vanitha Ramakrishnan | Maria Amelia Escobedo | Larry J. Fretto |
| Date | Date 6 28 635 | Date |

(Page 2 of 2)

K-112419-18-208C-2



DECLARATION AND POWER OF ATTORNEY

| As a below named inventor, I declare that: |
|--|
| My residence, post office address and citizenship are as stated below next to my name; I believe I am the original, first and sole inventor |
| (if only one name is besed below) or an original, first and joint inventor (if plural inventors are named below) of the subject matter which |
| is claimed and for which a patent is sought on the invention entitled: INHIBITORY IMMUNOGLOBULIN POLYPEPTIDES TO |
| HUMAN PDGF RETA RECEPTOR the specification of which is attached hereto or _X was filed on June 10, 1994 as |
| ADMAN I DOT META RECEIT TOX |
| Application Serial No. 08/258,283 and was amended on (if applicable). |

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| Country | Application No. | Date of Filing | Priority Claimed Under 35 USC 119 |
|---------|-----------------|----------------|--------------------------------------|
| | | | Yes No |
| | | | Yes No |

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Shelley Precivale Eberle, Rag. No. 31,411 Karen B. Dow, Reg. No. 29,684 Randolph T. Apple, Rag. No. 36,429

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Karen B. Dow

TOWNSEND and TOWNSEND KHOURIE and CREW

Steuart Street Tower

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Karen B. Dow Reg. No. 29,684 (415) 326-2400, Ext. 217

| Full Name of Inventor 1 | Last Name Ramakrishnan | First Name Vanitha | Middle Name or Initial | |
|----------------------------|---|----------------------------------|---------------------------------|-------------------|
| Residence & | City Relmont | State/Foreign Country California | Country of Citizenship India | |
| Post Office | Post Office Address 2019 Arbor Avenue | City Belmont | State/Country California | Zip Code 94002 |
| Full Name of Inventor 2 | Last Name Escohedo | First Name Maria | Middle Name or Initial Amelia | |
| Residence & | City San Francisco | State/Foreign Country California | Country of Citizenship Chile | |
| Post Office | Post Office Address 455 Upper Terrace #3 | City San Francisco | State/Country California | Zip Code 94117 |
| Full Name of Inventor 3 | Last Name Fretto | First Name Larry | Middle Name or Initial J. | |
| Residence & Citizenship | City Reimont | State/Foreign Country California | Country of Citizenship U.S.A. | |
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| Signature of Inventor 1 | ar | Signature of Inventor 3 |
|-------------------------|-----------------------|-------------------------|
| Vanitha Ramakrishnan | Maria Amelia Escobedo | Larry J. Fretto |
| | Date | Date 6.22-85 |
| Date 22 JUNE 199512 | Date | |

(Page 2 of 2)

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

| In re application of: |) |
|---|-----------------------|
| Vanitha Ramakrishnan, et al. |) Examiner: P. Gambel |
| Serial No.: 08/258,283 |) Art Unit: 1806 |
| Filed: June 10, 1994 |) |
| For: INHIBITORY IMMUNOGLOBULIN POLYPEPTIDES TO HUMAN PDGF BETA RECEPTOR |))) _) |

Honorable Commissioner of Patents and Trademarks Washington, D.C. 20231

STATEMENT OF FACTS UNDER 37 C.F.R. 1.48(a)

Sir.

We, Vanitha Ramakrishnan, Maria Amelia Escobedo, Larry J. Fretto and Nathalie Lokker, hereby declare as follows:

- 1. That Vanitha Ramakrishnan, Maria Amelia Escobedo and Larry J. Fretto are the coinventors and co-applicants of patent application Serial No. 08/253, 440, filed on June 7, 1994 (the "Parent"), which is a file-wrapper-continuation of Serial No. 07/801,795, filed on December 2, 1991, describing immunoglobulin polypeptides that block binding of platelet- derived growth factor ("PDGF") to the platelet-derived growth factor receptor.
- 2. That the Parent describes an immunoglobulin polypeptide, referred to as the monoclonal antibody 2A1E2, and characterizes it as binding to the fifth domain of the extracellular domain of the PDGF receptor.
- 3. That the present application is a continuation-in-part of the Parent and characterizes 2A1E2 as binding to the second domain of the extracellular domain of the PDGF receptor, instead of the fifth domain as described in the Parent.

- 4. That Nathalie Lokker identified the second domain binding characteristics of 2A1E2. Based upon the fact that she provided additional data that was included in the present application and because claims were included to recite an immunoglobulin polypeptide the recognized an epitope in the second domain, Nathalie Lokker was named as an inventor at the time that the present application was filed.
- 5. It has now become apparent that Nathalie Lokker was incorrectly named as an inventor, through error and without any deceptive intention. Based upon our information and belief, the facts leading up to this discovery, are as follows:
- a) The issue of inventorship was raised by Examiner Gambel in the Office Action dated December 23, 1994. The Examiner took the position that since the Parent application described the same antibody that was being described in the present application, any differences in the epitope mapping (fifth vs. second Ig-like domain) only reflected a re-evaluation of the fine specificity of the same 2A1E2 antibody and did not change the inherent properties (i.e., structure and function) of the same deposited hybridoma (HB 10938). In light of this, the Examiner questioned why the inventive entities of the Parent and the present application were different.
- b) On June 1, 1995, in a telephonic interview between the Examiner and Applicants' representative Karen B. Dow, the Examiner more specifically questioned the inventorship status of Nathalie Lokker in view of the Examiner's assertion that the binding characteristics of the 2A1E2 antibody were inherent in the antibody, which was described in the Parent.
- c) On several occasions after this interview, Ms Dow discussed the inventorship issue with Applicants' attorneys, Carolyn R. Adler and Shelley P. Eberle, and it was determined that the binding characteristics of the monoclonal antibody 2A1E2 were inherent in the antibody, therefore that Nathalie Lokker was not an inventor of the subject matter as claimed in the present application.
- d) During the week of June 19, 1995, Ms. Eberle discussed this issue with us and we realized that an error in inventorship had been made and that it was appropriate to correct inventorship by the deletion of Nathalie Lokker as an inventor.
- 6. That Vanitha Ramakrishnan, Maria Amelia Escobedo and Larry J. Fretto are the coauthors of a publication entitled "A Novel Monoclonal Antibody Dependent on Domain 5 of the Platelet-Derived Growth Factor Beta Receptor Inhibits Ligand Binding and Receptor Activation", <u>Growth Factors</u>, (8):253-265 (1993); and that Joseph J. Seroogy, James E. Tomlinson and David L. Wolf are also named as co-authors on the publication. The contributions of the authors not named as inventors is as follows:
- a) David L. Wolf made the molecular biological construct and used it to transfect cells to produce the antigen (extracellular domain of the PDGF beta receptor), as described in column 1, lines 1-23, on page 255 of the publication. David L. Wolf also immunized rabbits to produce rabbit polyclonal serum, as described in column 1, lines 31-34 on page 255 of the publication.

- b) James E. Tomlinson characterized the antigen-expressing cell lines made by David L. Wolf, as described in column 1, lines 23-43 on page 255 of the publication. This antigen was later used as the immunogen for the production of the monoclonal antibodies. James E. Tomlinson also characterized the rabbit polyclonal antibodies made by David L. Wolf. These antibodies were used to determine the recombinant expression of the modified extracellular PDGF beta receptor by Western blot analysis, as described in column 1, lines 28-31 on page 255 of the publication.
- c) Joseph J. Seroogy performed the partial purification of the antigen (extracellular domain of the PDGF beta receptor) that was used to determine antibody titer in the ELISA, as described in column 2 on page 255 of the publication.
- d) It is our belief that co-authors Joseph J. Seroogy, James E. Tomlinson and David L. Wolf are not co-inventors of the claimed subject matter of the present application.

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| In re application of: |) |
|---|----------------------|
| Vanitha Ramakrishnan, et al. |) Examiner: P. Gambe |
| Serial No.: 08/258,283 |) Art Unit: 1806 |
| Filed: June 10, 1994 |) |
| For: INHIBITORY IMMUNOGLOBULIN POLYPEPTIDES TO HUMAN PDGF BETA RECEPTOR |))) _) |

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| Dated: | |
|----------------|-----------------------|
| | Vanitha Ramakrishnan |
| Dated: 6/28/95 | morina a associated |
| | Maria Amelia Escobedo |
| Dated: | · |
| | Larry J. Fretto |
| Dated: | |
| | Nathalie Lokker |